

ROTARY-DRIVE DISPENSER

Statement of Related Cases

This case claims priority of U.S. Provisional Patent Application Serial No. 60/441,757, filed January 21, 2003, and incorporated by reference herein.

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Field of the Invention

The present invention relates generally to liquid dispensers, and more particularly to a liquid dispenser that is capable of controllably dispensing a plurality of liquids into a plurality of receivers.

Background of the Invention

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A variety of industrial and research applications require that ingredients, typically liquid, be dispensed into vessels or other types of receivers. Representative applications include formulation chemistry, mixing of dyes, pharmaceuticals, clinical research, genetic research, and combinatorial chemistry, to name a few.

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It is desirable for a dispenser that is intended for these applications to be capable of:

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- dispensing one or more liquid ingredients into one or more receivers;
- dispensing very small quantities of liquid (*e.g.*, nano-liter to micro-liters);
- dispensing liquid ingredients rapidly and accurately; and
- monitoring the dispensing operation with real-time analysis and feedback.

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But few if any are. Rotary-type dispensers, wherein the dispensing elements or the receivers are moved in circular fashion, are well suited, at least in concept, to dispensing one or more ingredients onto or into one or more receivers. In fact, several prior-art dispensers have been developed that use rotary motion in conjunction with a dispensing operation.

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U.S. Pat. No. 6,192,945 B1 to Ford et al. discloses a fluid dispenser for an automated biological reaction system. The dispenser includes a circular reagent tray that supports reagent fluid dispensers. The reagent tray rotates to successively

position individual fluid dispensers into a reagent delivery position over a slide. Once positioned, reagent is dispensed onto the slide and then a subsequent fluid dispenser is rotated into position.

U.S. Pat. No. 6,202,895 B1 to Fox discloses an apparatus for dispensing
5 cosmetic foundation compositions. The apparatus includes a manually-rotatable carousel that receives a plurality of plunger-type dispensing elements. Various reagents (*e.g.*, hair coloring, *etc.*) are stored in the dispensing elements. In use, the carousel is rotated to position a particular dispensing element over a mixing container. A desired amount of reagent is next dispensed into the mixing container.
10 The carousel is then rotated to position another dispensing element over the mixing container, and so forth.

U.S. Pat. No. 6,338,820 B1 to Hubbard et al. discloses an apparatus for performing assays at reaction sites. The apparatus includes a rotatable substrate having radially-arrayed reaction sites. The apparatus further includes a rotary
15 stepper motor that controllably rotates the substrate. The apparatus also has a dual-function head that includes a fluid dispenser for delivering fluid to a reaction site, as well as a sensor for receiving a signal from the reaction site. The dual-function head is suspended for linear movement across the rotatable substrate.

While illustrative of the manner in which rotary motion can be used in
20 conjunction with a dispensing operation, the dispensers mentioned above are not capable of rapidly and accurately dispensing a plurality of ingredients into a plurality of receivers to create a plurality of formulations. In fact, few if any prior art dispensers, rotary or otherwise, are capable of providing this capability. Furthermore, it is very difficult to accurately dispense very small quantities of liquid
25 or to monitor and correct the dispensing operation in real time.

Summary of the Invention

A liquid dispenser that uses rotary motion to provide a capability for rapid and accurate dispensing of one or more ingredients into one or more receivers is disclosed.

30 Functionally, the dispenser includes a drive system, a dispensing system, and a system controller. The drive system creates a relative motion between dispensing elements (*e.g.*, nozzles, *etc.*) and underlying receivers (*e.g.*, vials, *etc.*) so that they align on a continuing basis. Fluid is dispensed from the dispensing elements into the

receivers. The system controller coordinates the operation of the drive system and the dispensing system.

The dispenser can be configured in various ways. In one configuration, the dispensing elements are moved by the drive system to align with stationary receivers. In an illustrative embodiment of this configuration, the dispenser includes at least one arm that is coupled to a drive shaft. The arm has at least one dispensing element, which is coupled to one or more reservoirs. The reservoirs contain the ingredient(s) that are to be dispensed by the dispensing elements. The reservoirs are disposed on a reservoir support structure, which, like the rotatable arm, is coupled to the drive shaft.

A drive element rotates the drive shaft, causing both the arm and the reservoir support structure to rotate. Under the control of the system controller, the dispensing elements deliver liquid ingredients into one or more receivers. The receivers are disposed on a receiver support structure that is located between the dispensing elements and the reservoir support structure.

In another configuration, the receiver support structure, and the receivers along with it, are moved by the drive system to align with stationary dispensing elements. In an illustrative embodiment of this configuration, the dispenser has coaxial shafts: an inner shaft, which does not rotate, and an outer shaft, which does. At least one arm having at least one dispensing element is attached to the top of the non-rotating inner shaft. The dispensing elements are coupled to one or more ingredient-containing reservoirs. A receiver support structure is coupled to the rotatable outer shaft. As the shaft is rotated, the receiver support structure and the receivers move beneath dispensing elements to receive liquid.

In yet another configuration, the dispenser is capable of moving either the dispensing elements to stationary receivers or, in another mode of operation, moving the receivers to stationary dispensing elements.

In some variations of any of the configurations summarized above, the receiver support structure is a platform that comprises a plurality of pie-shape segments. Each segment, which is advantageously independently removable from the platform, includes an opening that is sized to accommodate a receiver. In some additional variations, each segment includes a stirrer motor that drives a magnet, which, in turn, drives a stirrer that is disposed in the receiver. Additionally, each segment can be provided with heating and cooling capabilities.

In additional variations of the present dispenser, each receiver includes one or more sample ports through which liquid is withdrawn from and returned to the receiver. This withdrawal and return process creates mixing or emulsification action within the receiver. In yet further variations, each receiver is coupled to a respective analysis window, which receives liquid from the receiver. The analysis window is configured to be used in conjunction with one or more analytical stations. The analytical stations are capable of providing analyses of the liquid that is retained within the analysis window. Some of the analyses are advantageously performed in real time, so that the results can be fed back to the system controller to make corrections to the dispensing operation.

The system controller implements and coordinates the activities of dispensing system and drive system in accordance with a dispensing protocol. In accordance with one dispensing protocol, ingredients are incrementally dispensed from the dispensing elements as a series of pulses. In accordance with this dispensing protocol, the various ingredients are added quasi-continuously (*i.e.*, a near-continuous flow of an ingredient) and/or quasi-simultaneously (*i.e.*, all ingredients combined at substantially the same time) to the receivers. See applicant's co-pending patent application entitled "Method and Apparatus for Quasi-Continuous and Quasi-Simultaneous" filed on January 21, 2003 as serial no. 10/348,769 and incorporated by reference herein.

When implementing the quasi-continuous dispensing protocol in conjunction with the present dispenser, the dispensing elements (or receivers) are in continuous motion. This provides several benefits and, in fact, solves certain "classic" dispensing problems. In particular, to the extent that the ingredients are dispensed by the present dispenser in accordance with the quasi-continuous dispensing protocol:

- a common manifold error that otherwise arises when a single reservoir feeds multiple arms/dispensing elements is overcome;
- the quantity of liquid delivered to each receiver can be determined to a high degree of accuracy using simple techniques; and
- problems can be rapidly corrected during dispensing operations using real time analysis and feedback.

These and other variations of a dispenser in accordance with the illustrative embodiment of the present invention are illustrated in the Drawings and described further in the Detailed Description section of this specification.

Brief Description of the Drawings

5 **FIG. 1A** depicts, via block diagram, basic functional components of a dispenser in accordance with the illustrative embodiment of the present invention.

FIG. 1B depicts, via perspective view, a first illustrative embodiment of a dispenser in accordance with the present invention, wherein dispensing elements are moved to bring them into alignment with receivers.

10 **FIG. 1C** depicts a drive for driving the drive shaft depicted in FIG. 1B.

FIGs. 2A, 2B, 2C, 2D, and 2E depict, via top views, several configurations for the rotatable member depicted in FIG. 1B.

FIG. 3 depicts a side view of a second illustrative embodiment of a dispenser in accordance with the present invention, wherein receivers are moved to bring them
15 into alignment with receivers.

FIG. 4 depicts a side view of a variation of the dispenser depicted in FIG. 3.

FIG. 5 depicts a side view of a third illustrative embodiment of a dispenser in accordance with the present invention, wherein either the dispensing elements or the receivers are moved.

20 **FIGs. 6A, 6B, 6C, 6D, and 6E** depict variations of the dispensing element used in a dispenser in accordance with the present invention.

FIG. 7 depicts a receiver support structure that is has a plurality of individually-removable segments.

FIGs. 8A and 8B depicts a stirrer for use in conjunction with the receivers.

25 **FIGs. 9A, 9B, and 9C** depict variations of a system for withdrawing liquid from receivers.

FIG. 10 depicts an analysis window for use in conjunction with the systems depicted in FIGs. 9A-9C.

30 **FIG. 11** depicts an analytical station for use in conjunction with the illustrative embodiments of a dispenser in accordance with the present invention.

FIG. 12 depicts the analytical station shown in FIG. 11 incorporated into the dispenser depicted in FIG. 1B.

FIG. 13 depicts a simplified block diagram of a control system for use in conjunction with the illustrative embodiments of a dispenser in accordance with the present invention.

FIG. 14 depicts a top view of the dispenser of FIG. 1 to accompany Example I, providing a "snap shot" of the dispenser at the beginning of a dispensing cycle.

Detailed Description

Definition of terms and additional considerations:

"Liquid(s)" means material(s) that are liquid at the temperature of operation of the dispenser, materials that are liquefied by various physical processes, liquid suspensions (*e.g.*, material(s) that are suspended in a liquid carrier, *etc.*), slurries, even solids that have properties that allow them to "flow," (*e.g.*, fluidized solids, *etc.*). Thus, the term "liquid" includes solids that are "naturally" flowable or rendered flowable using appropriate operations (*e.g.*, processing, *etc.*) or appropriate conditions (*e.g.*, temperature, *etc.*), *etc.*

"Fluid" means gases, vapors, and liquids.

"Coupled" means that (coupled) elements cooperate, communicate, attach to, or otherwise influence or affect one another. For example, fluid can flow between (fluidically) coupled elements (*e.g.*, a reservoir and a dispensing element coupled by a conduit, *etc.*). Also, a force exerted by or experienced by a first of two (mechanically) coupled elements can affect the second element, whether or not the two elements are directly attached to one another.

Tasks and Subtasks. The operation of the illustrative embodiment is described in terms of tasks and subtasks, rather than steps. This is because, as will be clear to those skilled in the art, some of the described tasks and subtasks can be performed in a single step, while others cannot. Furthermore, the illustrative embodiment is more easily understood when it is described in terms of its constituent tasks and subtasks than if it were described, formalistically, in terms of "steps."

Elements described in terms of their function. Some elements of the illustrative apparatus are described functionally, or in terms of the tasks or subtask

that they carry out. As will be clear to those skilled in the art, these elements can be implemented using shared or dedicated hardware including, for example, hardware capable of executing software, such as a suitably-programmed, general purpose processor.

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FIG. 1A depicts a block diagram of the basic functional components of dispenser **100** in accordance with the illustrative embodiment of the present invention. Dispenser **100** comprises dispensing system **101a**, drive system **101b**, and system controller **101c**.

10 The implementation details of dispensing system **101a** and drive system **101b** can vary as a function of which of the element(s) — the dispensing elements (e.g., nozzles, etc.), the receivers (e.g., vessels, etc.), or both — is moved by drive system **101b**. First, embodiments of dispenser **100** are described wherein the dispenser is configured so that the dispensing elements are moved into alignment
15 with stationary receivers. Next, embodiments of dispenser **100** are described wherein the dispenser is configured so that the receivers are moved into alignment with stationary dispensing elements. Finally, embodiments of dispenser **100** are described wherein the dispenser is configured so that either the dispensing elements or the receivers can be moved (into alignment with the other).

20 After describing the illustrative embodiments of these three basic configurations of dispenser **100**, a description of various features and optional features that can be used with any of the dispenser configurations is provided. Finally, a description of the system controller **101c** is provided and the operation of dispenser **100** in accordance with a quasi-continuous dispensing protocol is
25 described.

Dispenser **100** Configured to Move
the Dispensing Elements into
Alignment with Stationary Receivers

Referring now to FIG. 1B, dispensing system **101a** includes reservoir(s) **128-1** and **128-2** (generally "reservoir(s)" **128**), liquid-transport system(s) **130-1** and **130-2** (generally, "liquid-transport system(s) **130**"), dispensing element(s) **112-1** and **112-2** (generally "dispensing element(s) **112**"), and receiver support structure **116**.

In the illustrative embodiment, each reservoir **128** is coupled to a respective dispensing element **112** via a respective liquid-transport system **130**. As a consequence of this connectivity, liquid ingredients that are stored in reservoirs **128** can be delivered to receivers (e.g., receivers **142-1**, etc.) that are disposed on receiver support structure **116**.

With reference to FIGs. 1B and 1C, drive system **101b** includes drive **103**, drive shaft **114**, reservoir support structure **124**, and rotatable member **102**. (For clarity, only drive **103** and drive shaft **114** are depicted in FIG. 1C.) The elements of the drive system are coupled in such a way that drive **103** is capable of causing reservoir support structure **124** and rotatable member **102** to rotate. Rotation of rotatable member **102** brings dispensing elements **112** into alignment with underlying receivers **142-1**, etc. (into which liquid is dispensed). While drive **103** depicted in FIG. 1C is a belt drive, in some variations, a direct drive arrangement is used.

Position sensor **1372** (see FIG. 13), such as a rotary encoder, is used in conjunction with drive **103** so that system controller **101c** "knows" the position of each dispensing element **112** relative to receivers **142**, to the extent that the dispensing elements have moved. It is within the capabilities of those skilled in the art to select and integrate position sensor **1372** into dispenser **100**.

With continuing reference to FIG. 1B, rotatable member **102** includes two arms **104-1** and **104-2** (generally "arm(s) **104**"). The rotatable member is coupled, near its mid-point **106**, to drive shaft **114**. Dispensing element **112-1** depends from arm **104-1** near end **108** and dispensing element **112-2** depends from arm **104-2** near end **110**. Further description of dispensing elements **112** is provided later in this specification.

Receiver support structure **116**, which is implemented as a "plate" or "platform" (hereinafter referred to as "receiver support platform **116**" or simply "platform **116**,") is disposed beneath arms **104**. Platform **116** "surrounds" drive shaft **114** such that the platform and drive shaft are concentric. To this end, platform **116** has centrally-located opening **118** through which drive shaft **114** passes.

In accordance with this embodiment, platform **116** is not coupled to drive shaft **114**. In other words, platform **116** is not movable by the drive shaft. Since, in this embodiment, platform **116** is not coupled to drive shaft **114**, it must be

supported in some other fashion. For example, in some embodiments, platform **116** is supported by several legs (not depicted for the sake of clarity) that engage its lower surface and are attached to base **140**.

Even though, in this embodiment, dispensing elements **112** are moved into alignment with receivers **142** (rather than vice versa), it is nevertheless
 5 advantageous to provide platform **116** with an independent rotational capability (as indicated by arrows **146** and **148**). As described later in this Specification, this rotational capability enables dispenser **100** to be used in conjunction with an analytical station.

10 With continuing reference to FIG. 1B, upper surface **120** of platform **116** receives one or more receivers **142** at near-perimeter region **122**. Only one such receiver, receiver **142-1**, is shown in FIG. 1B for the sake of clarity. The radial position of dispensing elements **112** along arms **104** is such that they overlie near-perimeter region **122**. Therefore, to the extent that an arm **104** is angularly aligned
 15 with a receiver **142**, the dispensing element **112** that depends from that arm overlies the receiver.

Reservoir support structure **124** underlies platform **116**. In the illustrative embodiment, reservoir support structure **124** is implemented as a plate or platform, hereinafter referred to as "reservoir support platform **124**" or simply "platform **124**."
 20 Upper surface **126** of platform **124** receives one or more reservoirs **128** — two of which (reservoirs **128-1** and **128-2**) — are depicted in FIG. 1B. Reservoirs **128** store the ingredients that are to be dispensed by dispensing elements **112**. The ingredients are advantageously stored in liquid form in reservoirs **128**.

It is understood that platform **116** supports receivers **142** and that platform
 25 **124** supports reservoirs **128**. This functionality can, of course, be accomplished by structures other than a platform. For example, in some embodiments, the receiver support and/or reservoir support are cage structures that attach, via arms, to drive shaft **114**, etc.

In the illustrative embodiment, each reservoir **128** is coupled to one
 30 dispensing element **112** via liquid-transport system **130** so that liquid can flow from the reservoir to the dispensing element. In FIG. 1B, for example, reservoir **128-1** is coupled to dispensing element **112-1** via liquid-transport system **130-1** and reservoir **128-2** is coupled to dispensing element **112-2** via liquid-transport system **130-2**.

Liquid-transport system **130-1** comprises conduit **132-1** (and a check valve, which is not shown), positive-displacement pump **134-1** and conduit **136-1**. Likewise, liquid-transport system **130-2** comprises conduit **132-2** (and a check valve, which is not shown), positive-displacement pump **134-2** and conduit **136-2**.

5 With respect to liquid-transport system **130-1**, pump **134-1** takes its suction from reservoir **128-1**. When actuated, pump **134-1** draws liquid ingredient **138-1** from reservoir **128-1** through conduit **132-1** and pumps it through conduit **136-1** to dispensing element **112-1**. Dispensing element **112-1** then dispenses liquid ingredients **138-1** into an underlying receiver **142**. Liquid-transport system **130-2**
10 functions in the same fashion in conjunction with reservoir **128-2** and dispensing element **112-2**. Other arrangements for transporting liquid from reservoirs **128** to dispensing elements **112**, as will occur to those skilled in the art, can suitably be used.

 In some variations of the illustrative embodiment, dispensing elements **112**
15 must be actuated independently of any requirement for actuating liquid-transport system **130**. In some other variations, however, dispensing elements **112** do not need to be actuated. That is, when liquid is delivered to a dispensing element, the liquid is dispensed without any need to actuate the dispensing element itself. This is described in further detail later in this Specification.

20 With regard to pump **134**, any of a variety of different types of pumps can suitably be used. One type of pumping system that is particularly well suited for this service is a positive displacement pump. Positive displacement pumps, which are well known in the art, can be microprocessor controlled and use a piston-, roller- or peristaltic-type pumping mechanism. In the illustrative embodiment, pump **134** is
25 depicted as having a piston-type pumping mechanism, a common implementation of which is a syringe.

 In some other variations of the illustrative embodiment, liquid-transport system **130** does not use a pump. Rather, in these other variations, reservoir **128** is pressurized, such as by a line that runs from a source of pressurized gas to the
30 reservoir. An in-line control valve is used to regulate pressure. In yet some additional variations, neither a pump nor a source of pressurized gas is used in conjunction with liquid-transport system **130**. In those variations, a special nozzle (described later in this Specification) is used that draws liquid from reservoirs **128** by inducing a partial vacuum in the nozzle.

Like rotatable member **102**, reservoir support platform **124** is coupled to drive shaft **114**. With reference to FIG. 1C, drive shaft **114** is coupled to drive **103**, which is disposed within base **140**. When actuated, drive **103** (e.g., a motor, turbine, etc.) rotates drive shaft **114**, such as in the direction indicated by arrow

5 **144**. The drive shaft, in turn, rotates rotatable member **102** and reservoir support platform **124** in the same direction (see, FIG. 1B). The rotatable member and reservoir support platform are advantageously driven at the same rate (or otherwise linked) to ensure that conduits **136** will not twist and fail, as would otherwise occur if there were a relative rotational motion between rotatable member **102** and platform

10 **124**. Base **140**, which is advantageously weighted or otherwise quite heavy, provides stability when arms **104** and platform **124** are rotating.

In FIG. 1B, dispenser **100** is depicted as having two arms **104**, two dispensing elements **112**, one receiver **142**, and two reservoirs **128**. It is to be understood that these details are provided by way of illustration, not limitation. In

15 particular, in a typical application, more than one receiver **142** will be used in conjunction with dispenser **100**. For example, as a function of the size of platform **116** and the size of receiver(s) **142**, eight, twelve, sixteen, twenty four, or some other number of receivers **142** can suitably be used.

Furthermore, in the illustrative embodiment depicted in FIG. 1B, there are

20 two dispensing elements **112-1** and **112-2** and two reservoirs **128-1** and **128-2**. Reservoir **128-1** is coupled to dispensing element **112-1** and reservoir **128-2** is coupled to dispensing element **112-2**. In some variations of the illustrative embodiment, a single reservoir **128** is coupled to more than one dispensing element **112** (e.g., reservoir **128-1** to dispensing elements **112-1** and **112-2**, etc.). This

25 type of arrangement can be problematic, however, since it presents the problem of "common manifold error" (i.e., common manifolds never deliver identical volumes of liquid to parallel lines or nozzles). This topic is addressed later in this Specification. In some other variations of illustrative dispenser **100**, more than one reservoir **128** is coupled to a single dispensing element **112** (e.g., reservoirs **128-1** and **128-2** to

30 dispensing element **112-2**, etc.).

In some variations of the illustrative embodiment, a fixed number of reservoirs **128** (e.g., five, etc.) is always present on platform **124**. In those variations, if some of the reservoirs are not required, then they are either not

coupled to a dispensing element **112** or the associated liquid-transport system **130** is not actuated, etc.

It is to be understood that while rotatable member **102** depicted in FIG. 1B comprises two arms **104-1** and **104-2**, in other variations, the rotatable member
 5 can have a different number of arms. FIGs. 2A through 2E depicts several such variations of rotatable member **102**. In particular, FIG. 2A depicts rotatable member **202a** having a single arm **104-1**. FIG. 2B depicts rotatable member **202b** having three arms **104-1**, **104-2**, and **104-3**. FIG. 2C depicts rotatable member **202c** having four arms **104-1**, **104-2**, **104-3**, and **104-4** and FIG. 2D depicts
 10 rotatable member **202d** having five arms **104-1**, **104-2**, **104-3**, **104-4**, and **104-5**. And FIG. 2E depicts rotatable member **202e** having six arms **104-1**, **104-2**, **104-3**, **104-4**, **104-5**, and **104-6**.

One dispensing element **112** is advantageously disposed toward the end of each arm **104** such that when the arm is angularly aligned with a receiver **142**, the
 15 dispensing element **112** overlies the receiver. In particular, rotatable member **202a** includes dispensing element **112-1**, rotatable member **202b** includes dispensing elements **112-1**, **112-2**, and **112-3**, rotatable member **202c** includes dispensing elements **112-2**, **112-2**, **112-3**, and **112-4**, rotatable member **202d** includes dispensing elements **112-2**, **112-2**, **112-3**, **112-4**, and **112-5** and rotatable
 20 member **202e** includes dispensing elements **112-2**, **112-2**, **112-3**, **112-4**, **112-5**, and **112-6**.

The position of dispensing elements **112** near the ends of arms **104** is a consequence of situating receivers **142** in near-perimeter region **122**. In some other embodiments, receivers **142** are disposed inward of near-perimeter region
 25 **122** and, consistent therewith, dispensing elements **112** are disposed inward of the end of arms **104** so that they overlie the receivers. In other words, dispensing elements **112** must be radially aligned with receivers **142**.

Using a rotatable element **102**, **202** having a relatively greater number of arms **104** offers certain advantages. In particular, a group of receivers **142** can be
 30 filled more rapidly if more dispensing elements **112** are used. Also, the use of a relatively greater number of dispensing elements **112** provides an ability to individually dispense a relatively greater number of ingredients. In some variations (not shown) of the illustrative embodiment, at least two dispensing elements **112** are disposed along an arm **104**, one inward of the other. Positioning two or more

dispensing elements along arm **104** enables the arm to service multiple angularly-aligned receivers that are disposed at different radial locations.

In some variations of dispenser **100**, rotatable element **102, 202** is physically adapted to be readily removed from drive shaft **114**. This enables rotatable elements **102, 202** to be exchanged, *e.g.*, one having a relatively greater number of dispensing elements **112** for one having a relatively lesser number, *etc.*, as suits the needs of a particular application. In some other applications, dispenser **100** is fitted with a rotatable element **102, 202** that is typically not changed to satisfy application requirements. For example, in some embodiments, dispenser **100** is fitted with rotatable member **202e** having six arms **104** and six dispensing elements **112**. To the extent that some of dispensing elements **112** are not required for a particular application, they are simply not used (*e.g.*, liquid-transport system **130** for the particular dispensing element **112** is not actuated, *etc.*).

Dispenser **100** Configured to Move the Receivers into Alignment with Stationary Dispensing Elements

FIGs. 3 and 4 depict illustrative embodiments of dispenser **100** in which the dispenser is configured to move receivers **142** into alignment with stationary dispensing elements **112**. Both embodiments use a coaxial arrangement of shafts. In the variation depicted in FIG. 3, non-rotatable member **350** is attached to the top of inner shaft **352**. In FIG. 3, non-rotatable member **350** is configured like rotatable member **102** with one or more arms **104-1** and **104-2**. Inner shaft **352**, which does not rotate, conducts conduits **136-1** and **136-2** to respective arms **104-1** and **104-2**. These conduits couple reservoirs **128** to dispensing elements **112**.

Outer shaft **114** functions as a drive shaft, and is coupled to drive **103**. Receiver support platform **116** is attached to (or integral with) drive shaft **114**. When drive **103** is actuated, it turns drive shaft **114**, which causes platform **116** to rotate. As platform **116** rotates, receivers **142** move into position beneath dispensing elements **112** to receive liquid. Position sensor **1374** (see FIG. 13), such as a rotary encoder, is used in conjunction with drive **103** so that system controller **101c** “knows” the position of each receiver **142** relative to dispensing elements **112**, to the extent that platform **116** has moved. It is within the capabilities of those skilled in the art to select and integrate position sensor **1374** into dispenser **100**.

Since it is not necessary to rotate reservoirs **128** (unlike embodiments in which arms **104** are rotating) they can be placed directly on base **140**. Liquid is conducted from reservoirs **128**, through the appropriate conduits, pumps (not shown), *etc.*, and is dispensed through dispensing elements **112**.

5 In dispenser **100** depicted in FIG. 4, reservoirs **128** are disposed on top of non-rotating member **350**. In this variation, non-rotating member **350** is configured as a platform so that it can support reservoirs **128**. Dispensing elements **112** depend from the underside of non-rotating member **350**. Liquid can be gravity fed from reservoirs **128-1** and **128-2**, via respective conduits **454-1** and **454-2**, to
10 respective dispensing elements **112-1** and **112-2**. When actuated, drive **103** causes drive shaft **114** and receiver support platform **116** to rotate.

Dispensing elements **112** that depend from non-movable member **102** can be supplemented, as desired, by additional controlled dispensing elements (not depicted) that are supported by support arms (not depicted) that are attached, for
15 example, to base **140**.

Dispenser **100** Configured to Move Either the Receivers or the Dispensing Elements

FIG. 5 depicts a variation of dispenser **100** wherein receiver support platform **116** on the one hand, and arms **104** and reservoir support platform **124** on the
20 other hand, are capable of rotating independently of each other. In this variation, arms **104** and platform **124** are arranged and driven as depicted in FIG. 1b. Platform **116** is driven by a second drive; in this embodiment: receiver-support drive **556**. In the embodiment depicted in FIG. 5, drive **556** comprises motor **558** and belt **560**. The belt engages collar **562**, which is attached to platform **116**.
25 Collar **562** rides on bearing **564**. Motor **558** drives belt **560**, which, in turn, drives collar **562** and attached platform **116**. When a belt-type drive is used, as in the illustrative embodiment, collar **562** advantageously includes belt guides **724** (see FIG. 7).

Drive **556**, the lower portion of collar **562**, and bearing **564** are disposed in
30 housing **566**. The housing is supported by leg **568**, which is secured to base **140**. Position sensor **1374** (see FIG. 13), such as a rotary encoder, is used in conjunction with drive **556** so that system controller **101c** "knows" the position of each receiver **142** relative to dispensing elements **112**, to the extent that platform **116** has

moved. It is within the capabilities of those skilled in the art to select and integrate position sensor **1374** into dispenser **100**.

Dispensing Elements **112**

A variety of different types of dispensing elements **112**, some of which are depicted in FIGs. 6A through 6E, can be used in conjunction with the various configurations of dispenser **100** that have been described above. The particular type of dispensing element **112** that is used depends, to a certain extent, on the particular type of liquid-transport system **130** being used. For example, when liquid-transport system **130** incorporates a pump, such as a positive-displacement pump, dispensing element **112** can be simply embodied as an orifice, such as the open end **670** of conduit **136** (see, FIG. 1B). The orifice must be appropriately sized so that the liquid can be dispensed as a directed stream. The size of the orifice is a function of pressure/flow rate of the liquid that is being dispensed. When dispensing micro-liter liquid volumes, the size of the orifice is typically in a range of about 0.1 millimeters to 1 millimeter in diameter.

In a further variation of dispensing element **112** that can be used when liquid-transport system **130** includes a pump, dispensing element **112** comprises nozzle **672**, which is coupled to end **670** of conduit **136**, as depicted in FIG. 6B. The nozzle provides an orifice **674** that can be, for example, smaller in diameter than open end **670** of conduit **136**.

FIG. 6C depicts a further variation of dispensing element **112**, wherein dispensing valve **676** is coupled to end **670** of conduit **136**. The variation depicted in FIG. 6C can be used whether liquid-transport system **130** incorporates a pump or not (e.g., incorporates a pressurized reservoir instead of a pump, etc.). Suitable dispensing valves **676** are available from a large number of commercial suppliers. Such valves can be designed to handle milliliter, micro-liter or nano-liter volumes of liquid that is delivered in stream, droplet or aerosol patterns. It is understood that when dispensing element **112** is a valve, the valve must be actuated (in addition to any requirement for actuating liquid-transport system **130**) to dispense liquid.

Dispensing valve **676** advantageously possesses at least some of the following characteristics:

- very fast (about 1 millisecond response time) and repeatable opening and closing times; and

- has a very low CV (coefficient of variation) of about 5 percent or less; and
- an exit orifice that is shaped and dimensioned to reduce flow turbulence and reduce the incidence of satellite droplets; and
- 5 • the valve should have a minimum dead volume and otherwise possess shape and surface characteristics that decrease any tendencies for liquid losses and accumulation; and
- special materials or coatings to improve inertness of internal components and increase life expectancy; and
- 10 • a hard-on-hard valve seat, such as a sapphire to ruby valve seat.

These characteristics are particularly important when a very small volume (e.g., micro-liters or nano-liters) of liquid is being dispensed. A valve that possesses all of these characteristics is commercially available from TechElan of Mountainside, New Jersey ("Sub-micro-liter dispensing valve").

- 15 FIG. 6D depicts yet an additional variation of dispensing element **112**. The variation depicted in FIG. 6D is a "dual-flow" nozzle **678** that includes inner passageway **680** and surrounding shroud **682**. Liquid ingredient **138** is delivered via conduit **136**, under the control of an in-line valve (not depicted), to inner passageway **680** and dispensed through orifice **686**. An inert, secondary fluid **689**
- 20 (preferably but not necessarily a gas) is delivered, via conduit **684**, to shroud **682** and dispensed through annular opening **688**.

- Secondary fluid **689**, which is controlled independently of ingredient(s) **138**, forms a "sheath" that, at suitable rates of flow, reduces the tendency of dispensed liquid to "fan" or spread. The sheath focuses, concentrates or otherwise directs the
- 25 flow of liquid ingredient **138** toward a receiver. Furthermore, secondary fluid **689** is capable of accelerating the flow of liquid from nozzle **686** to a receiver, thereby speeding the dispensing operation. Additionally, secondary fluid **689** carries (into a receiver) any residual droplet that might otherwise form at orifice **686** at the completion of a dispense.

- 30 A further variation of dispensing element **112** is depicted in FIG. 6E. The variation depicted in FIG. 6E is a nozzle **690** that is similar to the dual-flow nozzle shown in FIG. 6D in that it includes an inner passageway **691** and a surrounding shroud **692**. In the variation that is depicted in FIG. 6E, however, inner passageway

691 is configured in the well-known venturi configuration. In particular, inner passageway **691** includes convergent region **693**, throat **694**, and divergent region **695**. Convergent region **693** typically has an included angle of about 20 degrees and divergent region **695** has an included angle of about 10 degrees.

Carrying fluid **698** (typically, but not necessarily, a gas) flows through inner passageway **691**. If ingredient(s) **138** are dispensed pulse-wise (see discussion of quasi-continuous dispensing protocol later in this specification), then the flow of carrying fluid **698** is, more accurately, a pulse of carrying fluid. A pulse of carrying fluid can be generated, for example, using a fast-acting pneumatic valve. Valves suitable for this service are available from FESTO, Inc (Germany) and other suppliers.

The flow of carrying fluid **698** through inner passageway **691** creates low pressure in throat **694**. The low pressure in the throat draws in liquid **138** that is "waiting" in conduit **136** (held there by capillary forces). Liquid ingredient **138** enters throat **694** through orifice **696** and is dispensed through orifice **697**. As a consequence of the venturi configuration, neither a pump nor a source of pressurized gas for pressurizing reservoir **128** is required as part of liquid-transport system **130**.

In some variations, independently-controlled, inert, secondary fluid **689** (preferably but not necessarily a gas) is delivered by a conduit (not shown) to shroud **692** and dispensed through annular opening **699**. As described above, secondary fluid **689** forms a "sheath" for focusing, *etc.*, the flow of liquid ingredient **138** from orifice **697**. In some embodiments, carrying fluid **698** and secondary fluid **689** are the same fluid.

It is noteworthy that carrying fluid **698** and secondary fluid **689** can reduce contact between the liquid within each receiver **142** and the ambient environment by blanketing each receiver **142**. Alternatively, one of dispensing elements **112** can simply be dedicated to dispensing inert gas to blanket receivers **142**.

Segmented Receiver Support Platform **116**

In some variations of the illustrative embodiment, receiver support platform **116** is composed of a plurality of pie-shaped segments **700**, as depicted in FIG. 7. This segmented platform can be used in conjunction with the various configurations of dispenser **100** that have been described above.

Each segment **700** (only one of which is shown for clarity) advantageously includes opening **702** for accommodating receiver **142**. Segment **700** also advantageously includes stirrer drive system **704**. In the embodiment depicted in FIG. 7, stirrer drive system **704** comprises motor **706**, belt **708**, and magnet **710**.
 5 Stirrer **712** having magnetic tip **714** is disposed within receiver **142**. Stirrer **712** will move to follow magnet **710** as it is driven by motor **706**. Unlike conventional stirrers that normal reside at the bottom of a vessel, stirrer **712** is elongated, like a swizzle stick. For relatively high-aspect ratio (*i.e.*, length to diameter) vessels, the elongated structure of stirrer **712** is expected to be more effective for stirring the
 10 contents of receivers **142**. Stirrer **712** is advantageously bent, as depicted in FIG. 8A, or has other adaptations for creating turbulence in the liquid contained in receiver **142**. For example, stirrer **712** depicted in FIG. 8B has "vanes" **818**. The increased turbulence generally results in improved mixing action.

Each segment **700** is advantageously removably attached to collar **562** at
 15 region **720**. Holes **722** in region **720** receive pins **716** of segments **700** in mating electrical registration. Power is distributed to each segment **700** via this connection. Power is supplied to collar **562** in known fashion, such as by using a slip-ring connector (not depicted). The power that is delivered to segments **700** can be used, for example, to operate stirrer drive **704** and to energize heating or cooling elements
 20 **726**. The heating/cooling elements **726** can be electrically-based (*e.g.*, resistive, infrared for heating or Peltier for cooling) and advantageously use a thermal conduction means (not shown), such as liquids, springy metal sponges, thermally-conductive beads, and the like to promote heat transfer to or from receiver **142**.

The size and shape of segments **700** advantageously vary to accommodate
 25 different-size vessels. And to that end, segments **700** are preferably sized as multiples of a minimum segment size. For example, a minimum-size segment can be 15 degrees, with other larger segments as multiples thereof (*e.g.*, 30 degrees, 45 degrees, *etc.*).

In FIG. 7, segment **700** is depicted as mating with collar **562** at region **720**.
 30 This collar is used in the embodiment of dispenser **100** that appears in FIG. 5. It will be understood that drive shaft **114** of FIGs. 3 and 4 can be physically adapted (*e.g.*, incorporate region **720**, see FIG. 7) for mating with segments **700**.

Sampling/Mixing System 925

In some variations of the illustrative embodiment, dispenser **100** includes sampling/mixing system **925**, illustrative examples of which are depicted in FIGs. 9A – 9C and 10. Using sampling/mixing system **925**, a portion of the contents of a receiver **142** can be removed and then returned to the receiver. Each receiver **142** requires its own sampling/mixing system **925**.

When used with sampling/mixing system **925**, receivers **142** are advantageously provided with one or more ports **926**, as depicted in FIGs. 9A through 9C. Port(s) **926** connect to conduit **928**, which is coupled to aspirator/dispenser mechanism **934** (e.g., a syringe, bellows, etc.). Aspirator/dispenser mechanism **934** is actuated by actuator **935**.

In operation, a sample of liquid is withdrawn from receiver **142** through port(s) **926** and then returned to the receiver through the same port(s) via the action of aspirator/dispenser mechanism **934**. This creates a mixing action within receiver **142**. When two ports **926** are used, as in depicted in FIGs. 9B and 9C, a capability for intense agitation or emulsification is provided. In FIG. 9B, two ports **926** are oriented at an angle with respect to one another (e.g., between 1 to 179 degrees, etc.), and, in FIG. 9C, the two ports **926** are diametrically opposed (i.e., 180 degrees) thereby generating active, turbulent, counter-flow mixing. End portion **930** of conduit **928** couples to one of the ports, while end portion **932** couples to the other. Liquid that is returned through the two ports co-mingles and emulsifies.

In some variations, sampling/mixing system **925** incorporates analysis window **1036**, depicted in FIG. 10. Analysis window **1036** provides a widened region in which liquid from a reservoir **142** is distributed for optical analysis, such as with analytical station **1142** described below.

As depicted in FIG. 10, conduit **926** couples to port **1038a** of analysis window **1036**. Conduit **1040** is coupled, at one end, to port **1038b** and at the other end to aspirator/dispenser mechanism **934**. Liquid drawn from reservoir **142** by aspirator/dispenser mechanism **934** is distributed throughout analysis window **1036**. On the return stroke, etc., of aspirator/dispenser mechanism **934**, the liquid within analysis window **1036** is returned to reservoir **142**. Analysis window **1036** can be formed, for example, from two, spaced-apart quartz plates that are attached to a frame. The gap between the plates for retaining liquid is in a range of about 1mm, with a total volume between plates of 5 to 50 micro liters.

Analytical Station **1142**

It is often desirable, during the course of dispensing operations, to sample the contents of receivers **142** for compositional analysis. The results of the analysis can be used for a variety of purposes. For example, samples can be taken at regular intervals to monitor reactions and/or obtain kinetic data. Another use for analytical data is to close a control loop that monitors the composition of the mixture or its properties.

Using analytical equipment to monitor formulation operations presents certain challenges or difficulties. One difficulty is that any on-going agitation or heating/cooling processes typically have to be interrupted in order to obtain a sample. Furthermore, if it is desirable to sample plural receivers **142** simultaneously, then each receiver requires an independent analytical system. And if sampling is performed sequentially with a single sampling system, then the system must be cleaned between each sampling operation. A further consideration is that, for many uses of analytical data, the analysis must be performed very rapidly after sampling to be of any value. Consequently, the analytical equipment should be located near to the dispenser. But situating analytical equipment near to the dispenser is not always practical since the typically small amount of available bench-top real estate might be required for other types of equipment.

In some variations of the illustrative embodiment, dispenser **100** incorporates integrated analytical station **1142**. One embodiment of integrated analytical station **1142**, which addresses the issues described above, is depicted in FIGs. 11 and 12.

With reference to FIG. 11, analytical station **1142** includes emitter **1144** and detector **1146**. In the illustrative embodiment, optional plates **1148** and **1150** are disposed between the emitter and detector defining a testing region **1152** therebetween. In some embodiments, plates **1148** and **1150** are used as a guide to conduct or route analysis window **1036** into testing region **1152**. In such embodiments, plates **1148** and **1150** are advantageously optically transparent (at least at the wavelengths of interest). In some other embodiments, plates **1148** and **1150** provide a filtering function by filtering out certain wavelengths from the radiation that is being delivered by emitter **1144** or that is being received by detector **1146**. The various elements (e.g., emitter **1144**, etc.) of analytical station **1142** are disposed on support **1154**.

In operation, analysis window **1036**, containing liquid from reservoir **142**, is moved into testing region **1152** between emitter **1144** and detector **1146**. Radiation having a wavelength that is appropriate for the analysis being conducted is emitted from emitter **1144**. The emitted radiation passes through the liquid sample in analysis window **1036**. Radiation that is transmitted through analysis window **1036**, or that is generated (*e.g.*, fluorescence, *etc.*) when the sample is exposed to the radiation, is detected at detector **1146**. Signal **1156**, which is generated by detector **1146** in response to the received radiation, is received by appropriate analysis electronics (not depicted).

The analysis electronics is capable of processing the signal, in known fashion, for determining certain properties, characteristics, *etc.*, of the sample. Typical analytical tests include spectrophotometry (at wavelengths ranging from ultraviolet to infrared), light scattering detection techniques, differential refractive index detection, capillary-viscometric detection, to name but a few.

FIG. 12 depicts one way in which analytical station **1142** can be integrated with dispenser **100**. In this embodiment, analytical station **1142** is integrated with dispenser **100** having movable arms **104** (*see, e.g.*, FIG. 1B). It is to be understood that analytical station **1142** can be integrated with any of the configurations of dispenser **100** described herein.

As depicted via the "cut-away" view provided in FIG. 12, analytical station **1142** is disposed beneath receiver support platform **116**. Analytical station **1142** is supported by support arm **1258**, which is attached to base **140**. Support arm **1258** suspends analytical station **1142** near the underside of receiver support platform **116** in position to receive analysis window **1036**.

To engage analytical station **1142**, analysis window **1036** is disposed beneath platform **116**. To that end, conduits **926** and **1040** pass through receiver support platform **116**.

In operation, liquid is drawn (aspirated) from reservoir **142** into analysis window **1036** by aspirator/dispenser **934** in conjunction with actuator **935**. Analysis window **1036** is rotated into position in testing region **1152** (*see* FIG. 11) by rotating receiver-support platform **116** (*i.e.*, in the illustrative embodiment, receiver-support platform **116** is rotated clockwise in direction **146**). To do so, dispenser **100** must incorporate a means for rotating receiver support platform **116**, such as receiver support drive **556** (*see*, FIG. 5 and the accompanying description). For this

service, drive **556** operates at slow speed, and, at most, rotates platform **116** about one full revolution. Alternatively, drive **556** can provide discontinuous motion. That is, platform **116** is rotated until analysis window **1036** reaches analytical station **1142** and then stops. This is in contrast to the continuous motion of platform **116** that occurs when operating in accordance with the quasi-continuous dispensing protocol (in configurations of dispenser **100** wherein receivers **142** are rotated). Consequently, a different type of drive **556** might be selected in embodiments in which the drive is used to move receiver support platform **116** for the dispensing operation (and for movement in conjunction with analytical station **1142**) as compared to those embodiments in which drive **556** is used only in conjunction with analytical station **1142**. In fact, in some embodiments in which receiver support platform **116** is used for both services, two different drives are coupled to platform **116**, one for each service. It is within the capabilities of those skilled in the art to select a drive suitable for either one or both of these services.

After the analysis is completed, liquid in analysis window **1036** is pumped, via aspirator/dispenser **934**, back into receiver **142**. Analysis window **1036** is rotated out of testing region **1152** and another analysis window that is associated with another receiver **142** is rotated into the testing region.

Although only one analytical station **1142** is depicted in FIG. 12, in some variations, a plurality of analytical stations are integrated with dispenser **100**. In some embodiments, each of the analytical stations performs the same test, enabling rapid testing of the samples in the plurality of receivers **142**. In some other embodiments, the various analytical stations perform different tests.

System Controller **101c**

The operation of dispensing system **101a**, drive system **101b**, sampling/mixing system **925**, and analytical station **1142** are coordinated by system controller **101c**. These operations are conducted in accordance with any one of a variety of dispensing protocols (some are described later in this specification), as implemented via system controller **101c**.

FIG. 13 depicts a block diagram of the salient components of system controller **101c**. Among other capabilities, the system controller is capable, in some embodiments, of:

- causing drive system **101b** to create a relative motion between dispensing elements **112** and receivers **142**;
- causing dispensing system **101a** to dispense ingredients;
- monitoring the dispensing operation; and
- adjusting the dispensing operation.

As will be understood by those skilled in the art, some of the components that compose system controller **101c** are implemented using shared or dedicated hardware including, for example, hardware capable of executing software, such as a suitably-programmed, general purpose processor, *etc.*

In the embodiment depicted in FIG. 13, system controller **101c** includes processor **1360**, memory **1362**, drive-system controller **1364**, drive-system drivers **1368**, position sensors **1372** and **1374**, dispensing-system controller **1366**, dispensing-system drivers **1370**, flow sensors **1376**, and analytical **1378**, interrelated as shown.

Processor **1360** is a general-purpose processor that is advantageously capable of performing the tasks described below. In particular, and without limitation, processor **1360** is capable of:

- receiving data from a local input device (not shown), such as a keyboard, control panel, touch display, *etc.*, and outputting it to a local output device (not shown), such as a monitor, signaling panel, *etc.*;
- storing parameters that are required to determine the rate at which drive shaft **114** is driven and when and for how long liquid is dispensed into any particular receiver **142**, and retrieving those parameters from memory **1362**;
- executing one or more software programs that are stored in memory **1362**;
- storing, in memory **1362**, the results of any software programs that have been executed; and
- controlling the operation of drive-system controller **1364** and dispensing-system controller **1366**.

Memory **1362** is a non-volatile memory (*e.g.*, an EEPROM, disk drives, an optical device, *etc.*) for storing:

- programs **1380** that are executed by processor **1360**;
- input parameters **1382** that are required for executing programs **1380**;
- properties data **1384** (*e.g.*, physical properties/pressure/flow data) for use with some of programs **1380**; and
- results **1386** that are generated by processor **1360**, among other information.

Drive-system controller **1364** converts commands (*e.g.*, move at a certain speed, *etc.*) that are issued by processor **1360** into drive- (*e.g.*, motor-) control instructions. Drivers **1368** carry out the drive-control instructions thereby causing drive **103** to operate. As a consequence, drive shaft **114** is rotated. In a dispenser that is configured like dispenser **100** depicted in FIG. 1B, rotation of drive shaft **114** causes rotatable member **102** to rotate. In a dispenser that is configured like dispenser **100** depicted in FIG. 3, rotation of drive shaft **114** causes receiver support plate **116** to rotate.

As previously indicated, drive-system controller **1364** moves rotatable member **102** (or receiver support platform **116**) so that dispensing elements **112** align, on a continuing basis, with receivers **142**. Position sensor(s) **1372** provides positional information to processor **1360** (or drive-system controller **1364** depending upon design specifics) so that the processor “knows” the position of dispensing elements **112** relative to any given receiver **142**. In some other embodiments, an encoder (absolute or incremental) is used for the same purpose.

Dispensing-system controller **1366** converts commands (*e.g.*, dispense now, *etc.*) that are issued by processor **1360** into actuator- (flow-) control instructions. Drivers **1370** carry out the flow-control instructions thereby causing liquid to be dispensed. For example, in the illustrative embodiment depicted in FIG. 1B, controller **1366** and drivers **1370** cause pump(s) **134** to activate and pump liquid from reservoir(s) **128** to dispensing element(s) **112**.

Optional flow measurement data, as provided by flow sensor **1376** can be used to close a control loop, so that the flow of ingredients is appropriately adjusted.

Obtaining the flow rate of ingredients when dispenser **100** is operating in accordance with the quasi-continuous dispensing protocol is described later in this specification.

Optional analytical measurement data (*e.g.*, composition, *etc.*), as obtained from analytical **1378** can be fed back to processor **1360** to adjust dispenser operation as appropriate. By way of example, emitter **1144** is advantageously coupled, through appropriate drivers and controllers (not depicted), to processor **1360**. The processor coordinates the movement of receiver support platform **116**, the actuation of aspirator/pump **934** and the activation of emitter **1090**. Similarly, detector **1146** is coupled to processor **1360** so that output signal **1156** (see, FIG. 11) from the detector, or from various analysis electronics (not depicted) that are associated with the detector, is routed to the processor. Processor **1360** uses the results of the analyses to adjust, as required, aspects (*e.g.*, flow rate of a particular ingredient, *etc.*) of the dispensing operation. To do this, process **1360** sends a signal to dispensing system controller **1366** to take the required action, which, in conjunction with drivers **1370**, is carried out.

Those skilled in the art will know how to design and implement the various components and software composing system controller **101c**.

Operating Dispenser **100** in Accordance with a Quasi-Continuous Dispensing Protocol

Dispenser **100** can be operated in accordance with any of a variety of dispensing protocols, three of which are described below.

In accordance with a first, well-known protocol, the full amount of a first liquid ingredient that is to be dispensed into a first receiver **142-1** is so dispensed, and then the full amount of the first ingredient that is to be dispensed into a second receiver **142-2** is so dispensed. Dispensing of the first ingredient continues, one receiver at a time, until each receiver (that is to receive the first ingredient) receives its portion. Then, the full amount of a second liquid ingredient that is to be dispensed into one or more of the receivers **142** is so dispensed, one receiver at a time. Dispensing continues, one ingredient at a time and one receiver at a time, until all ingredients are dispensed in accordance with the formulation.

In accordance with a second, well-known protocol, the full amount of a first liquid ingredient and the full amount of a second liquid that are to be dispensed into a first receiver **142-1** are so dispensed, simultaneously. Simultaneous dispensing,

receiver-by-receiver, of the first and second ingredient, in the full amount, continues until each receiver (that is to receive the ingredients) receives its portion.

In these well-known protocols, drive system **101b** moves (dispensing elements **112** or receivers **142**) in a *stepped* or discontinuous fashion.

5 Alternatively, in accordance with a quasi-continuous dispensing protocol, liquid ingredients are dispensed as a plurality of pulses. Each pulse typically contains a small portion of a total quantity of an ingredient to be delivered to a receiver. For some formulation applications, each pulse contains an amount of ingredient within a range of between ten percent to about twenty percent of the total amount of the
10 ingredient to be delivered to any one receiver **142**. Representative applications include those wherein the full quantity of the required ingredients must be delivered very rapidly to receivers **142**.

For most formulation applications, each pulse contains an amount of ingredient that falls into one of two ranges. In a first range, each pulse contains an
15 amount of ingredient within a range of about one percent to about ten percent of the total amount of the ingredient to be dispensed to any one receiver **142**. In the second range, each pulse contains an amount of ingredient within a range of about one-tenth of one percent to about one percent of the total amount of the ingredient to be dispensed to any one receiver **142**.

20 In yet some further formulation applications, each pulse contains less than one-tenth of one percent of the total amount of an ingredient to be delivered to any one receiver **142**. Given a total quantity of liquid to be dispensed, the smaller the quantity of the ingredient that is contained in each pulse, the longer it will take to dispense the full quantity of the ingredient.

25 Furthermore, in accordance with the quasi-continuous dispensing protocol, a single pulse of a first liquid ingredient is received by substantially all receivers before any receiver gets a second pulse of the first liquid. Yet, this second pulse typically occurs within 0.5 to about 5 seconds after the first pulse. Consequently, the first liquid ingredient is dispensed in an almost-continuous (*i.e.*, quasi-continuous)
30 fashion to each receiver.

In some variations of the quasi-continuous dispensing protocol, all receivers get a pulse of all other ingredients that are to be dispensed before any receiver gets a second pulse of the first liquid. Thus, the ingredients are dispensed in an almost-

simultaneous (*i.e.*, quasi-simultaneous) fashion on both an intra- and inter- receiver basis. That is, a given receiver receives one pulse of each of the required ingredients at substantially the same time, and the various receivers receive ingredients at substantially the same time. In some variations, of the quasi-continuous dispensing protocol, the dispensing is both quasi-continuous and quasi-simultaneous.

When operating dispenser **100** in accordance with the quasi-continuous dispensing protocol, drive system **101b** moves (dispensing elements **112** or receivers **142**) in *continuous* fashion.

10 The quasi-continuous dispensing protocol has certain benefits or advantages over known dispensing protocols. These benefits, which are summarized below, are described in greater detail in "Method and Apparatus for Quasi-Continuous and Quasi-Simultaneous Dispensing," U.S. Pat. Application serial number 10/348,769.

15 One benefit of the quasi-continuous dispensing protocol is its ability to serve multiple vessels with one dispensing mechanism per liquid. This increases the reliability of a dispenser that is operating in accordance with the protocol (relative to a combinatorial-type dispensing system).

20 Another benefit of the quasi-continuous dispensing protocol is an ability to dispense a near-constant proportional addition of multiple liquid ingredients into multiple receivers. This capability is crucial for the preparation of time-based, reactive formulations (*e.g.*, polymer formulations using different monomers and additives, *etc.*).

25 When the quasi-continuous dispensing protocol is implemented using rotary-drive dispenser **100**, additional benefits or advantages are obtained over other protocols or other types of dispensers.

30 One such advantage is that the flow rates of individual ingredients, which are quite low, can be determined to a high degree of accuracy. Normally, when dispensing very small amounts of liquid at high speed, it is difficult to accurately determine the rate at which liquid is dispensed. Rather than measuring the flow from each dispensing element, the total or integrated flow of ingredient from each reservoir **128** is measured (to a high degree of accuracy) using trivial methods (*e.g.*, a flow meter, *etc.*). And, knowing the timing parameters for the delivery of each ingredient into each receiver **142** (*i.e.*, the time duration of a pulse of a particular

ingredient into a particular receiver), the flow into each receiver is obtained by a simple apportioning calculation. This statistically-averaged flow (the flow from each dispensing element is not actually measured) provides a very accurate and true "total dispensed volume."

5 A second advantage of operating dispenser **100** in accordance with the quasi-continuous dispensing protocol is that any operational problems (*i.e.*, clogged nozzle, *etc.*) will be readily detected as a deviation in expected flow from a reservoir **128**. Since ingredients are dispensed pulse-wise in small increments to most or all receivers, no one receiver should be affected to a greater degree than any other
10 receiver by such operational problems.

A third advantage of operating dispenser **100** in accordance with the quasi-continuous dispensing protocol is that operational problems such as a partially occluded nozzle can be readily corrected by simply increasing pulse length or pressure. Since no one receiver will be affected to any greater degree than another
15 receiver, the dispensing operation can continue without endangering the overall formulation and mixing process.

A fourth advantage of operating dispenser **100** in accordance with the quasi-continuous dispensing protocol is that it overcomes the common manifold problem. In particular, to the extent that a single reservoir **128** is feeding two or more
20 dispensing elements **112**, any inconsistencies in the amount of ingredient that is delivered to each dispensing element (the amounts will not be equal) will be averaged over all receivers **142**. That is, assuming each receiver **142** gets a pulse of a first ingredient from each of two dispensing elements **112**, it does not matter that the amounts of the first ingredient delivered from the dispensing elements are
25 not equal.

Rotary-drive dispenser **100**, when operating in accordance with the quasi-continuous dispensing protocol, therefore eliminates most of errors in combinatorial formulations or any proportional fillings.

Dispenser **100** operating in accordance with a quasi-continuous dispensing
30 protocol incorporates:

- **"Means for determining flow rate,"** which is defined for use in this description and the appended claims as instrumentalities for measuring bulk flow from a reservoir and for apportioning said flow on a per ingredient, per receiver basis. Means for determining flow rate includes, in

some embodiments, a flow meter for measuring bulk flow from each reservoir and a processor **1360** running software **1380** for operating in accordance with a quasi-continuous dispensing protocol.

- **“Means for real-time adjustment of flow rate,”** which is defined for use in this description and the appended claims as instrumentalities for changing the quantity of an ingredient contained in a pulse. The quantity of an ingredient in a pulse can be changed by changing the length (*i.e.*, time) of a pulse, or the operating pressure. Means for real-time adjustment includes, in some embodiments, processor **1360** running software **1380** for operating in accordance with a quasi-continuous dispensing protocol, dispensing system controller **1366**, and drivers **1370**.
- **“Means for distributed dispensing,”** which overcomes the common manifold problem, is defined for use in this description and the appended claims as follows. Means for distributed dispensing includes processor **1360** running software **1380** for operating in accordance with a quasi-continuous dispensing protocol, and a liquid delivery system that delivers a first pulse of an ingredient to each of a plurality of receivers before delivering a second pulse of the ingredient to any of the receivers.

EXAMPLE I

Tables I and II below provide an illustration of the operation of dispenser **100** depicted in FIG. 1B operating in accordance with a quasi-continuous dispensing protocol. For the illustration, dispenser **100** includes six reservoirs **128-1** through **128-6** (not depicted for clarity) and rotatable member **102** having six arms **104-1** through **104-6** and six dispensing elements **112-1** through **112-6**. For this example, only five of the reservoirs (*i.e.*, reservoirs **128-1**, **128-2** and **128-4** through **128-6**) are dispensing respective ingredients **A**, **B**, **C**, **D**, and **E** through five dispensing elements (*i.e.*, dispensing elements **112-1**, **112-2** and **112-4** through **112-6**). Ingredients are to be dispensed into eight receivers **142-1** through **142-8**. See, FIG. 14. The cycle time (*i.e.*, the time it takes for all the receivers to receive a small amount of each of the formulation ingredients via a “pulse” from each dispensing element) is 0.8 seconds. Each of the eight receivers receives the same amount — 10 milliliters — of ingredients **B**, **C**, **D**, and **E**; however, the amount of ingredient **A** that is dispensed to each of receivers **142-1** through **142-8** is varied. In particular, while receiver **142-1** receives a total of 10 ml of ingredient **A**, the other receivers receive a reduced amount as follows (in milliliters):

Receiver:	142-1	142-2	142-3	142-4	142-5	142-6	142-7	142-8
Quantity:	10.0	9.8	9.5	9.2	9.0	8.8	8.6	8.4

In some embodiments, the deficit in total liquid volume that results from the reduction in an ingredient (e.g., ingredient A, etc.) can be compensated for by the addition of an appropriate amount of an inert liquid compound. Alternatively, if the difference in total volume of the various formulations is relatively small, then it is preferable not to add any diluent to compensate for lost volume. Such decisions are best made on a case-by-case basis by those skilled in the art. For the present example, no additional liquid is added.

Thirty minutes is allotted to produce the formulations. Since each dispensing cycle takes 0.8 seconds, a total of: (30×60) seconds \times 1 cycle per 0.8 seconds = 2250 cycles are required.

Assuming that ingredients **B**, **C**, **D**, and **E** are dispensed in equal (quantity) pulses, each pulse of ingredients **B**, **C**, **D**, and **E** contains: $10 \text{ ml} / 2250 \text{ cycles} = 4.44 \text{ micro-liters of liquid}$.

As to ingredient **A**, the quantity of liquid in the pulses varies as a function of which receiver receives the ingredient. Assuming that 2250 equal pulses are delivered to any given receiver, the pulses contain the following quantities (in micro-liters) of ingredient **A**:

Receiver:	142-1	142-2	142-3	142-4	142-5	142-6	142-7	142-8
Quantity:	4.44	4.36	4.22	4.09	4.00	3.91	3.82	3.73

Tables I and II below show the first and second dispensing cycles, respectively, for the Example. FIG. 14 shows dispenser **100** (top view) as each cycle begins (1st pulse), wherein:

dispensing element **112-1** dispenses ingredient **A** into receiver **142-1**;
 dispensing element **112-2** dispenses ingredient **B** into receiver **142-3**;
 dispensing element **112-3** does not dispense an ingredient (into receiver **142-4**);

dispensing element **112-4** dispenses ingredient **C** into receiver **142-5**;
 dispensing element **112-5** dispenses ingredient **D** into receiver **142-7**; and
 dispensing element **112-6** dispenses ingredient **E** into receiver **142-8**.

TIME, sec	0.0	0.1	0.2	0.3	0.4	0.5	0.6	0.7
RECEIVER	1 ST PULSE	2 ND PULSE	3 RD PULSE	4 TH PULSE	5 TH PULSE	6 TH PULSE	7 TH PULSE	8 TH PULSE
142-1	4.44A	4.44E	4.44D	—	4.44C	—	4.44B	—
142-2	—	4.36A	4.44E	4.44D	—	4.44E	—	4.44B
142-3	4.44B	—	4.22A	4.44C	4.44D	—	4.44C	—
142-4	—	4.44C	—	4.09A	4.44E	4.44E	—	4.44C
142-5	4.44C	—	4.44B	—	4.00A	4.44E	4.44D	—
142-6	—	4.44C	—	4.44B	—	3.91A	4.44E	4.44D
142-7	4.44D	—	4.44C	—	4.44B	—	3.82A	4.44E
142-8	4.44E	4.44D	—	4.44C	—	4.44B	—	3.73A

Table I: First Dispensing Cycle – Quantity of Ingredient Dispensed (μ l)

With reference to Table I and FIG. 14, after the first pulse of the first cycle has occurred, 4.44 micro-liters of ingredient **A** has been dispensed into receiver **142-1**, 4.44 micro-liters of ingredient **B** has been dispensed into receiver **142-3**, 4.44 micro-liters of ingredient **C** has been dispensed into receiver **142-5**, 4.44 micro-liters of ingredient **D** has been dispensed into receiver **142-7**, and 4.44 micro-liters of ingredient **E** has been dispensed into receiver **142-8**. During this first pulse, no ingredients are dispensed into receivers **142-2**, **142-4**, and **142-6**, since no dispensing element **112** was near to receivers **142-2** and **142-6** and dispensing element **112-3**, which is aligned with **142-2**, is not dispensing any ingredient for this particular formulation.

At 0.1 seconds, the second pulse begins. By the time the second pulse begins, the rotatable member **102** has rotated counterclockwise. Dispensing element **112-1** is now in position to dispense ingredient **A** into receiver **142-2**, and so forth. In accordance with the formulation specifics, only 4.36 micro-liters of ingredient **A** are delivered to receiver **142-2**.

After the eighth pulse of the first cycle, 4.44 micro-liters of ingredients **B**, **C**, **D**, and **E** have been delivered to each of receivers **142-1** through **142-8**. And an amount of ingredient **A** between 4.44 micro-liters (into receiver **142-1**) and 3.73

micro-liters (into receiver **142-8**) is dispensed into the receivers **142** in accordance with the protocol.

- By the end of the seventh pulse (elapsed time of 0.7 seconds), a first pulse of all ingredients **A** through **E** are received, for example, by receiver **142-1**. This illustrates "quasi-simultaneous" dispensing in accordance with the dispensing protocol. At 0.9 seconds, the first pulse of the second cycle begins, such that about 0.8 seconds elapses between successive deliveries of ingredient **A** into receiver **142-1**. This illustrates "quasi-continuous" flow.

TIME, sec	0.8	0.9	1.0	1.1	1.2	1.3	1.4	1.5
RECEIVER	1 ST PULSE	2 ND PULSE	3 RD PULSE	4 TH PULSE	5 TH PULSE	6 TH PULSE	7 TH PULSE	8 TH PULSE
142-1	8.88A	8.88E	8.88D	—	8.88C	—	8.88B	—
142-2	—	8.72A	8.88E	8.88D	—	8.88C	—	8.88B
142-3	8.88B	—	8.44A	8.88E	8.88D	—	8.88C	—
142-4	—	8.88B	—	8.18A	8.88E	8.88D	—	8.88C
142-5	8.88C	—	8.88B	—	8.00A	8.88E	8.88D	—
142-6	—	8.88C	—	8.88B	—	7.82A	8.88E	8.88D
142-7	8.88D	—	8.88C	—	8.88B	—	7.64A	8.88E
142-8	8.88E	8.88D	—	8.88C	—	8.88B	—	7.46A

Table II: Second Dispensing Cycle — Cumulative Ingredient Dispensed (μ l)

Table II records, on a pulse-by-pulse basis, the accumulation of ingredients in each of receivers **142-1** through **142-8**.

- From the forgoing, it should be understood that the above-described embodiments are merely illustrative of the present invention and that many variations of the above-described embodiments can be devised by those skilled in the art without departing from the scope of the invention). It is therefore intended that such variations be included within the scope of the following claims and their equivalents.